Joca et al. - Supplement 4

| Bias Category | Questions | Gilbert-Diamond et al., 2011 | Wei et al., 2013 |
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| Selection bias | Were the comparison groups appropriate? | Definitely low risk of bias Women were divided into two groups - "rice eaters" who reported rice consumption within two days of giving the sample, and "non-rice eaters" who did not report consuming rice within two days. The women recruited for this study were between the ages of 18-45, were all within 24 to 28 weeks of gestation, all reported using a private, unregulated water system for which arsenic levels were measured, and recorded their water, food, and rice consumption for three days leading up to providing a urine sample. Therefore, risk of bias from inappropriate comparison groups is low. | probability cluster sampling, |

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| Performance bias | Did deviations from the study protocol impact the results? | Definitely low risk of bias - no deviations from the study protocol | Definitely low risk of bias - no deviations from the study design |

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| Attrition/Exclusion Bias | Were demographic data incomplete due to attrition or exclusion from analysis? | Definitely low risk of bias - no data were excluded from reporting | Probably low risk of bias - exclusion was limited to individuals under 20 and participates with missing data or below the lower detection level for urinary arsenic species |

| Were the outcome assessors blinded to study group or exposure levels? | this study measured exposure to arsenic and rice consumption; so no health outcomes were examined; if considering the arsenic measurements in water/urine as outputs, then although not explicitly stated, it is unlikely | Probably low risk of bias - manuscript examined relationship between urinary arsenic concentrations, race, and rice consumption, so no health outcomes were measured; the basis of the analysis is NHANES data and although not explicitly stated, it's likely that analysis of urinary arsenic concentrations were performed blinded to demographics/dietary information |
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| anticipated to bias results? | Definitely low risk of bias - smoking, water consumption, seafood, rice consumption considered as sources of arsenic | Probably low risk of bias - covariates included fish/shellfish consumption, source of drinking water; smoking was not explicitly examined |
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Probably low risk of bias -Definitely high risk of bias authors used total arsenic and exposures were estimated based upon self-reported DMA, rather than inorganic intake of water, rice, rice arsenic species, for analyses; products, and fish/seafood dietary assessment based upon rather than having a duplicate recollection could also diet analysis; different strains introduce uncertainty of rice have different iAs levels, so using an average for the brown rice strains available unlikely to reflect actual exposure; postpartum questionnaire on brown vs white rice consumption could Can we be confident in the introduce error into the exposure characterization? exposure estimates; cooking water for rice not reported requires correction adjustment

| Can we be confident in the outcome assessment? methods used to measure arsenic species/concentration are appropriate MHANES used established methods to determine urinary arsenic levels and demographics information; dietary information collected using questionnaire could introduce uncertainty | | arsenic species/concentration | correlations within NHANES data and didn't collect health outcome data; likely that NHANES used established methods to determine urinary arsenic levels and demographics information; dietary information collected using questionnaire could |
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| | | Definitely low risk of bias - measured demographics were reported | |
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| Selective Reporting W Bias | ere all measured demographics reported? | | |
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| Balazs et al. 2012 | Diawara et al., 2006 | Landolt et al., 1985 | O'Rourke et al., 1999 |
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| Probably high risk of bias comparison groups were either treated CWS or untreated CWS, however, only 6 CWS had confirmed arsenic treatment plants - therefore, the "treated" CWS may not have reduced arsenic; also, selection criteria led to underrepresentation of CWS with <200 connections | random selection, no apparent effort to match with | exposures across demographics; frequency of fishing activites also varied | NHEXAS AZ study; population-based probability design used to select households; females over-represented, but ethnicity was consistent with census data so impact |

| comparison groups were either treated CWS or untreated CWS, however, | the soil samples were taken along 4 transects, transects based upon socioeconomic parameters rather than random selection, no apparent effort to match with areas of higher SES/lower | interviews of fisherman at 4 locations; anglers primarily Caucasian and U.S. born - discrepancy in sampling could introduce bias when comparing potential for exposures across demographics; frequency of fishing activites also varied | NHEXAS AZ study; population-based probability design used to select households; females over-represented, but ethnicity was consistent with census data so impact |
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| | there were no deviations from the study protocol | The authors begin their | Definitely low risk of bias - no deviations from the study protocol |

| Probably low risk of bias - only deviation was if data were not available in PICME database, then used the Water Quality Monitoriing database to calculate population size | Definitely low risk of bias - there were no deviations from the study protocol | Probably low risk of bias - The authors begin their interviews going in 6-hour block, 4 times a day. Well into their study, they change protocol by going at the most popular times to interview anglers. | Definitely low risk of bias - no deviations from the study protocol |
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| Definitely low risk of bias - demographics information was estimated based upon Census data and GIS information; there were no data exclusions | Probably low risk of bias - demographic information based upon US Census data, focus on minority and low SES | Definitely low risk of bias - no data were excluded from reporting | |

Probably low risk of bias -Probably low risk of bias -Probably low risk of bias -Probably low risk of bias study examined this study used soil samples measurements of arsenic in the study selected relationships between to measure arsenic fish tissue were households and the samples arsenic exposure, CWS, and concentrations; it is unclear if independent of the were collected and sent for demographics - there was those measurements were fishermen interviews analysis; although not no health outcomes made knowledge of the specifically stated in the measured or reported; sampling site; however, the manuscript, it's unlikely the considering the statistical impact of that knowledge on measurements were made the measurements is likely analysis, the demographics with knowledge of the and arsenic/CWS treatment minimal; also if the demographic information data were collected from "outcome" is the arsenic distribution, it's unlikely that separate sources prior knowledge would impact the GIS results

| The authors do not control for other exposures, but were seeking out MCL violations for arsenic, which used direct testing | Because the authors use chemical analyses to assess the concentrations of the certain metals, there is not a large chance for confounding | Definitely high risk only fish consumpt considered; no consideration of did differences, water consumption, or sm | ion etary | Probably low risk of bias - exposure from air, water, and food were evaluated; also used duplicate diet study to confirm diet diary; smoking status not |
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| for arsenic in the water. Because of the unlikelihood that another chemical would alter results for arsenic, the lack of control leaves no risk of bias. | variables to cause altered levels of the soil metals. | | | considered is limitation |
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Definitely high risk of bias - Definitely high risk of bias -Definitely high risk of bias - Probably low risk of bias exposure estimates were sampling was limited and non-only total arsenic measured; arsenic species were not based upon averages of the random, with some exposure is estimate based presented, although the source concentrations. oversampling of low SES technique for measuring upon consumption and fish which assumed equal and areas; exposure is limited to species caught; no direct arsenic could distinguish independent contribution to soil concentrations and does measures of exposure between species and the the exposure that was not consider other sources of authors acknowledge the constant over time; difference in toxicity arsenic; no arsenic speciation 'treated" CWS did not data; no measures of between organic and explicitly treat for arsenic; exposure in humans inorganic forms; exposure model does not incorporate arsenic exposures were not measured in individuals weighting factors for route of exposure or individual susceptiblity; no confirmation of arsenic exposure via biomarkers

| between CWS, arsenic exposure, and demographics and did not | Probably high risk of bias - the distribution of the arsenic relies on nearest neighbor, which would be impacted by the selective sampling approaches taken | exposure by comparing fish | exposure and demographics |
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| Probably low risk of bias - demographics information based upon census tract data and GIS analysis to estimate communities served by CWS, latino and non-white were combined; The authors group together all minority demographics into a single measure "people of color" including Latino and non-Latino people of color. | Probably high risk of bias - demographics were reported and based upon US Census data (i.e., self-identification); errors are unlikely to impact conclusions | Definitely low risk of bias - all demographics data collected were reported | Definitely low risk of bias - demographics information was reported |
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| Pellizzari et al., 1999 | Tyrrell et al., 2013 | Johnson et al., 2011 | Postma et al., 2011 |
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| Probably low risk of bias - households identified as part of NHEXAS Region V study; used a stratified, four- stage, probability based design; differences between sample and census data for | Definitely low risk of bias - data collected as part of NHANES study; 5 independent cross-sectional | Probably high risk of bias - age, race, sex, water sources, and potentially smoking status were different between the populations, suggesting the comparison groups were different | Probably high risk of bias - study looked at households served by non-regulated |
| | | | |

| Probably low risk of bias - households identified as part of NHEXAS Region V study; used a stratified, four stage, probability based design; differences between sample and census data for teenagers and income may introduce bias | data collected as part of NHANES study; 5 independent cross-sectional | age, race, sex, water sources, and potentially smoking status were different between the populations, suggesting the comparison groups were | served by non-regulated |
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| Definitely low risk of bias - no deviations from study protocol | the authors describe | Definitely low risk of bias - no deviations from study protocol | Definitely low risk of bias - no deviations from study protocol |

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| Definitely low risk of bias - demographic data was collected and reported | exclusion of subjects from the study was described; only Mexican hispanics, Non-hispanic whites, and non-hispanic blacks included in the analyses because of "very few non-Mexican hispanics and other ethnicity groups" in the data; although authors did not explicity state threshold for analysis | The analysis was ultimately based on who took and submitted adequate toenail clippings. N=88 in Appalachia, n=151 in | Definitely low risk of bias - the study was a sample, so no demographics information was excluded |

| Probably low risk of bias - outcome (i.e., the measures of arsenic) was assessed on samples collected from households; although not explicitly stated, it's unlikely those measuring the arsenic concentrations were aware of the demographic information | Probably low risk of bias - samples were collected as controls for another study; it's likely that measurements of As concentration in toenails were done without knowledge of the groups | Probably low risk of bias - study measured arsenic concentrations in tap water; although not explicitly stated, it's likely that measurements were done without knowledge of the household demographics |
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| Definitely low risk of bias - air, food, beverage, water, and dust sources were monitored for arsenic modiator but was not investigated further because there was no data in NHANES; drinking water levels of iAs differ geographically and may have different impacts in different locations; also other dietary sources of arsenic were not analyzed | controlled for in the comparison | Probably high risk of bias - dietary differences or smoking status not considered |
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| Probably low risk of bias - | Definitely high risk of bias - Definitely high risk of bias - Not Applicable - |
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| no arsenic speciation data; | blood and urine specimens no arsenic speciation; ~70% study only measured arsenic |
| no confirmation of arsenic | taken from random of samples were below level in water; there was no |
| exposure via biomarkers, | subgroups selected by of detection for As, measurement or estimate of |
| study doesn't provide | NHANES, but measures are differences in diet could exposure |
| details on sample | total arsenic (organic and contribute to toenail As and |
| collection/analysis, but as | inorganic combined) or diet was not considered as a |
| | arsenobetaine; no measure of source of arsenic |
| likely followed the | iAs |
| procedures outlined | |
| previously | |
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| | Probably high risk of bias - the association between arsenic exposure and demographics is limited because of the large number of arsenic samples that were below detection limits | Definitely high risk of bias methods to measure arsenic in water were not indicated |
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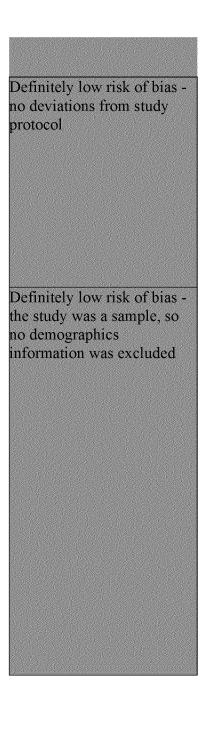
| Probably low risk of bias - demographics information was collected but not shown in study | The authors have | Definitely low risk of bias - all measured demographics were reported | Definitely low risk of bias - all demographics collected by interviewers were reported |
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Walker et al., 2005

Probably low risk of bias study looked at households
with private well water
consumption for potential
arsenic exposure;
households sampled in
approximately same pattern
and density as population;
age, homeownership, and
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Definitely low risk of bias no deviations from study protocol



Probably low risk of bias - study measured arsenic concentrations in tap water; although not explicitly stated, it's likely that measurements were done without knowledge of the household demographics

Probably high risk of bias -dietary differences or smoking status not considered

Definitely high risk of bias no arsenic speciation, exposure estimated by self-reported consumption of tap water (no biological measures of expsoure), no individual measures of exposure

Definitely low risk of bias measures of arsenic in water
had appropriate QA/QC
measures and were
performed using established
protocols

Probably high risk of bias -The authors do not report demograhic results, although they were collected in the methodology